

Mean # of lung metastases

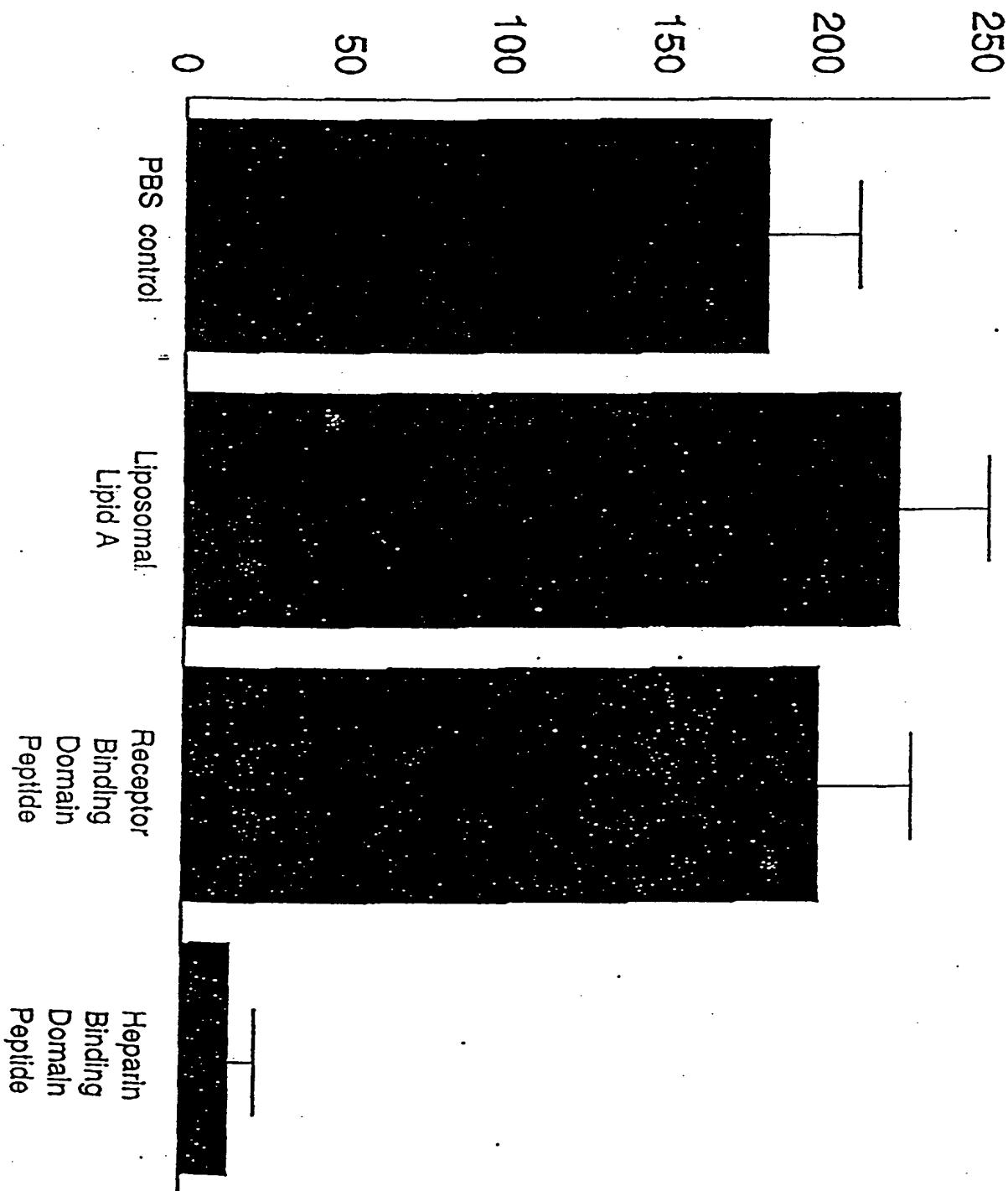


Figure 1

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Title: Compositions and Methods for Treating Cancer
Hyperproliferative Disorders

Inventors: John Holaday; Antonio Ruiz; John Madsen; Stacy Plum

Heparin Binding Domain Peptide

Liposomal Lipid A

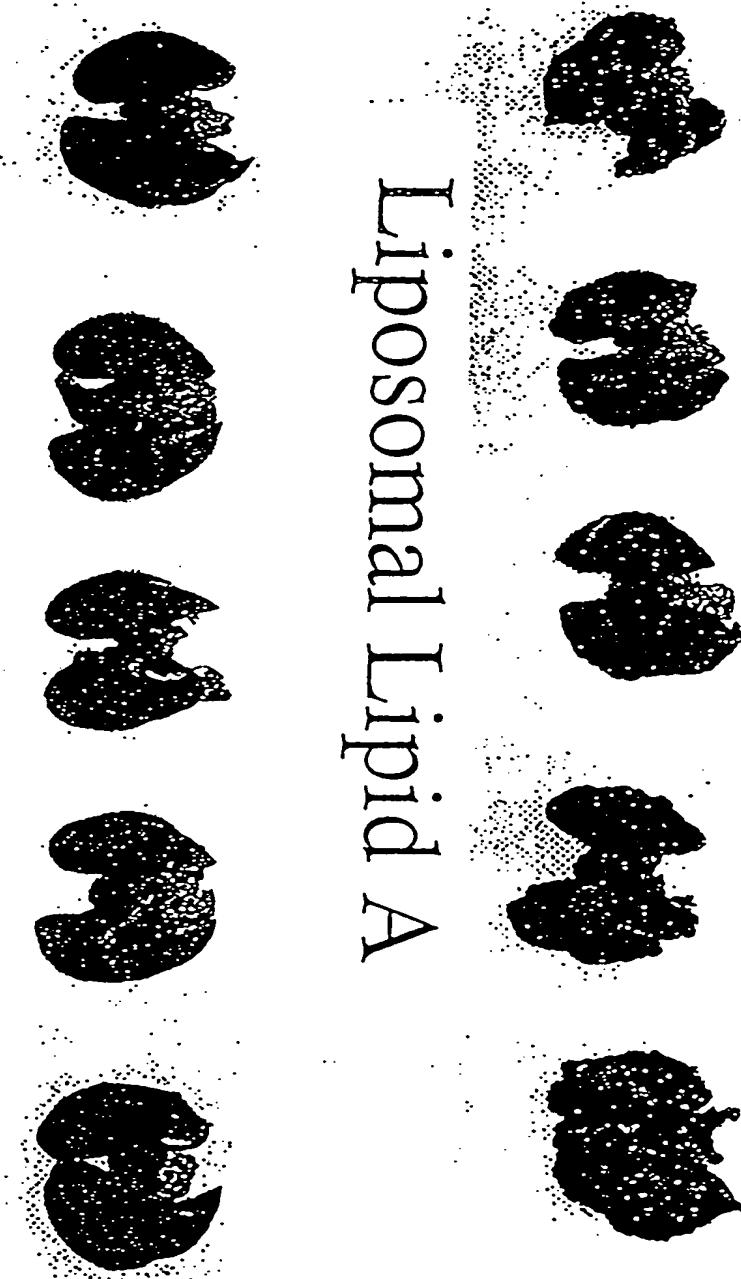
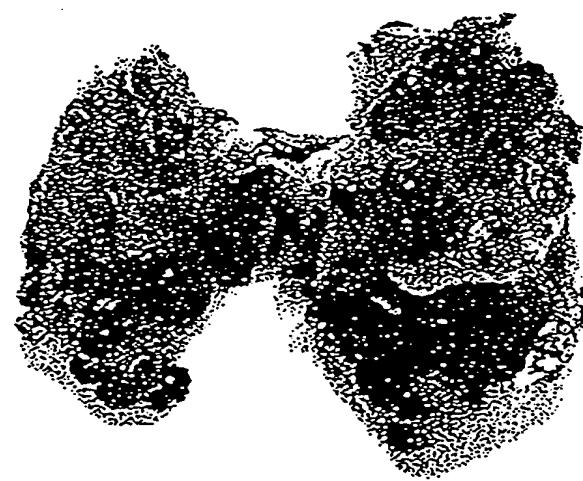
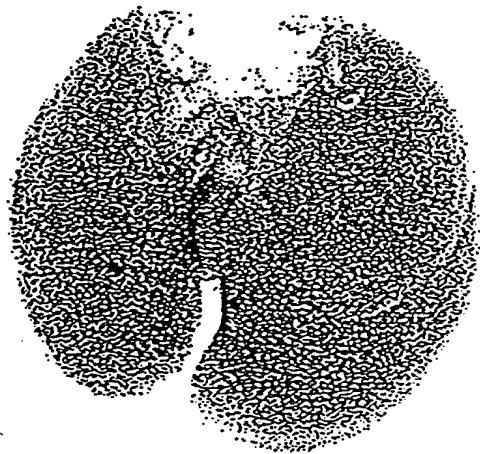


Figure 2

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HEPARIN BINDING DOMAIN

LLA CONTROL

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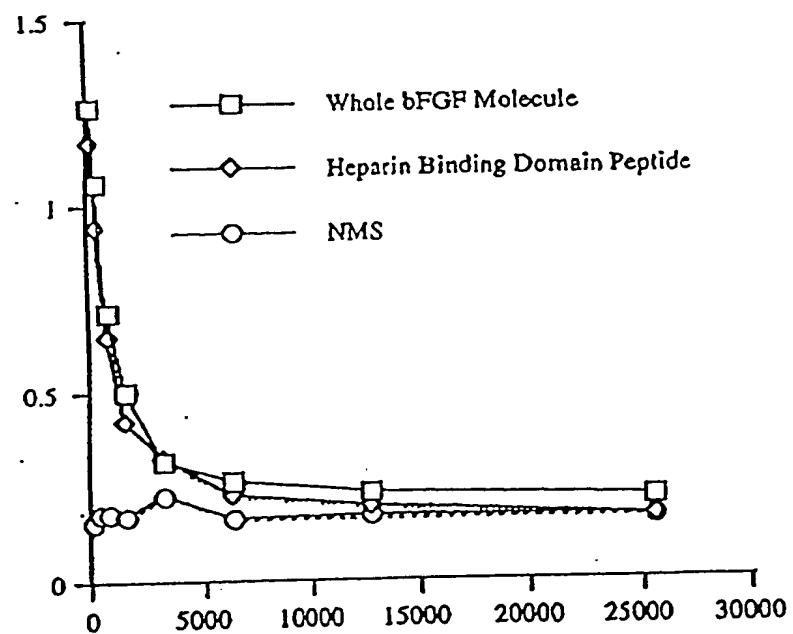


Figure 4

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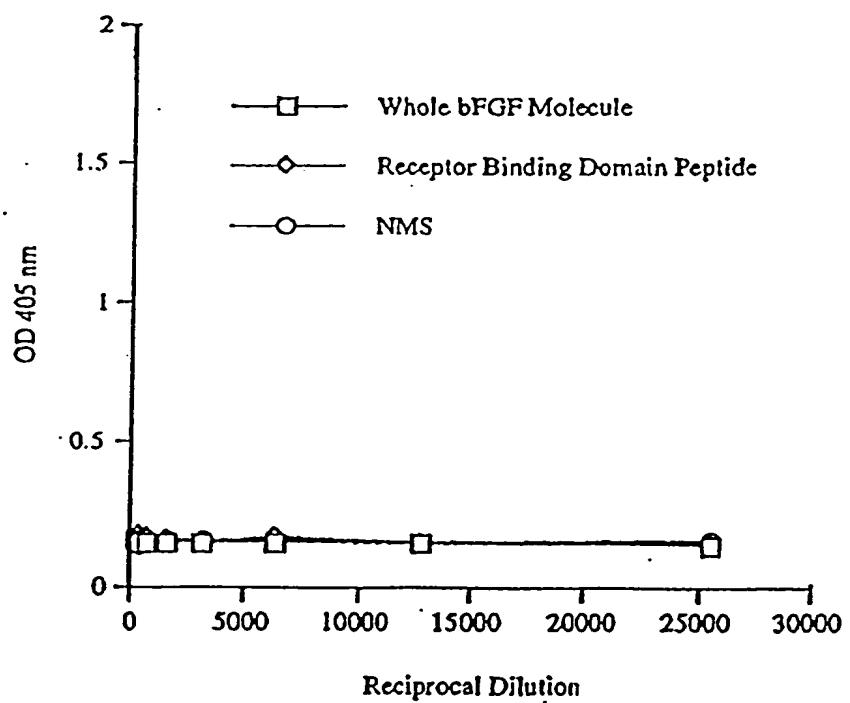


Figure 5

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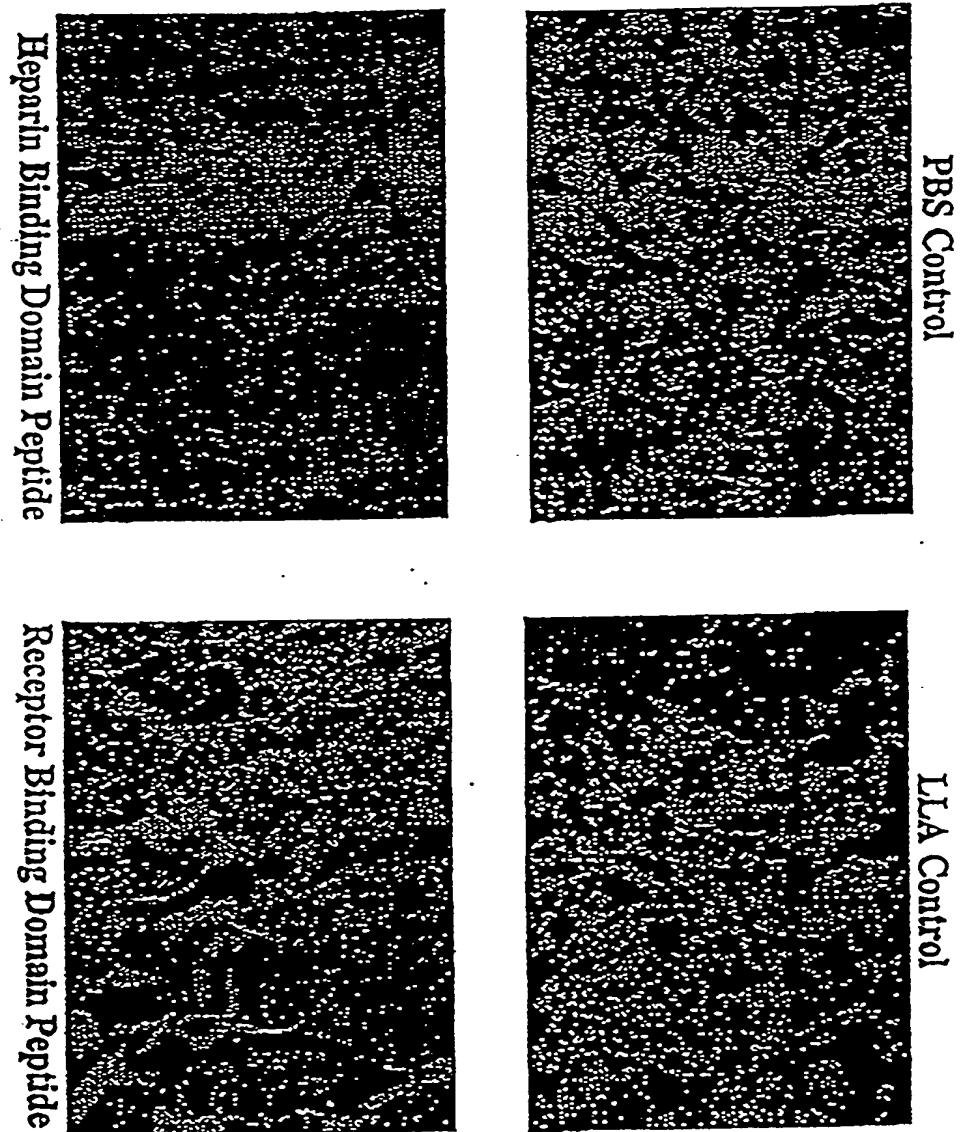


Figure 6

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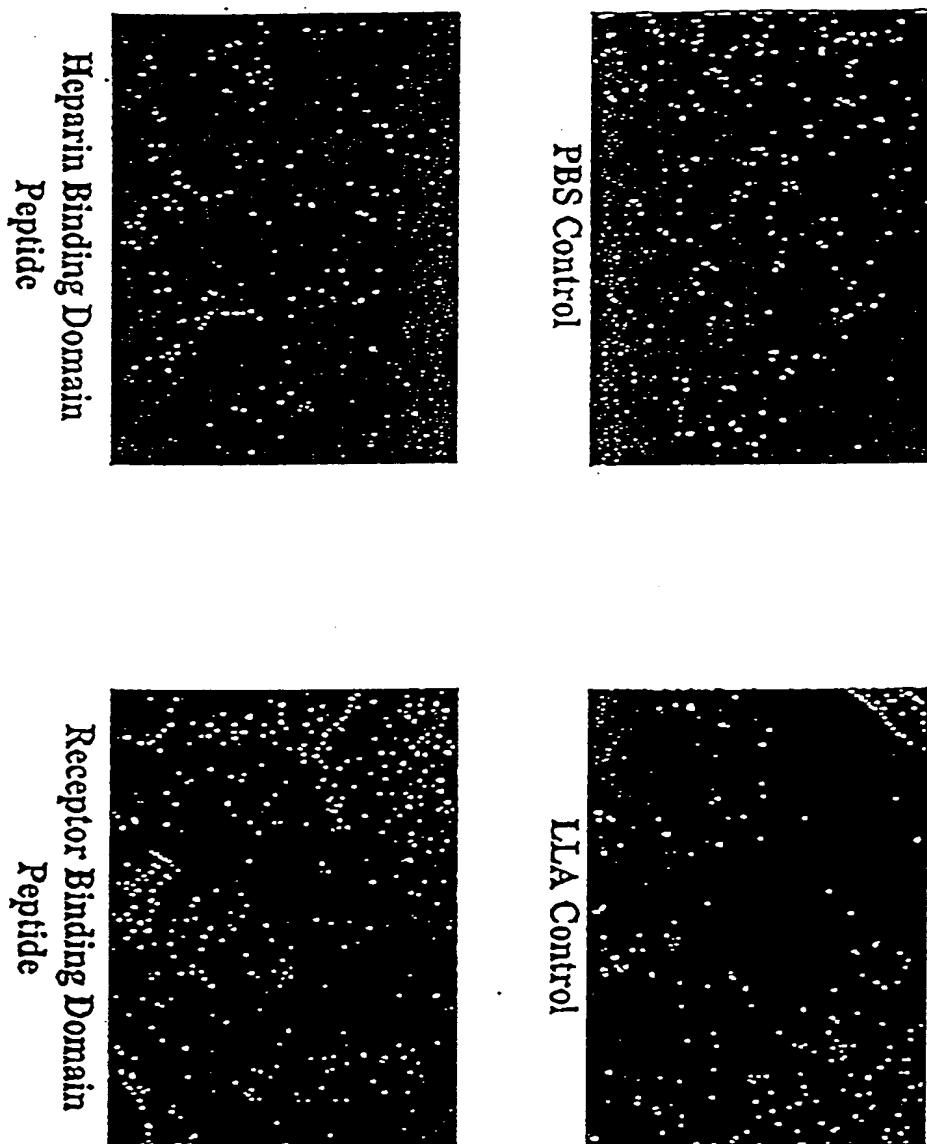


Figure 7

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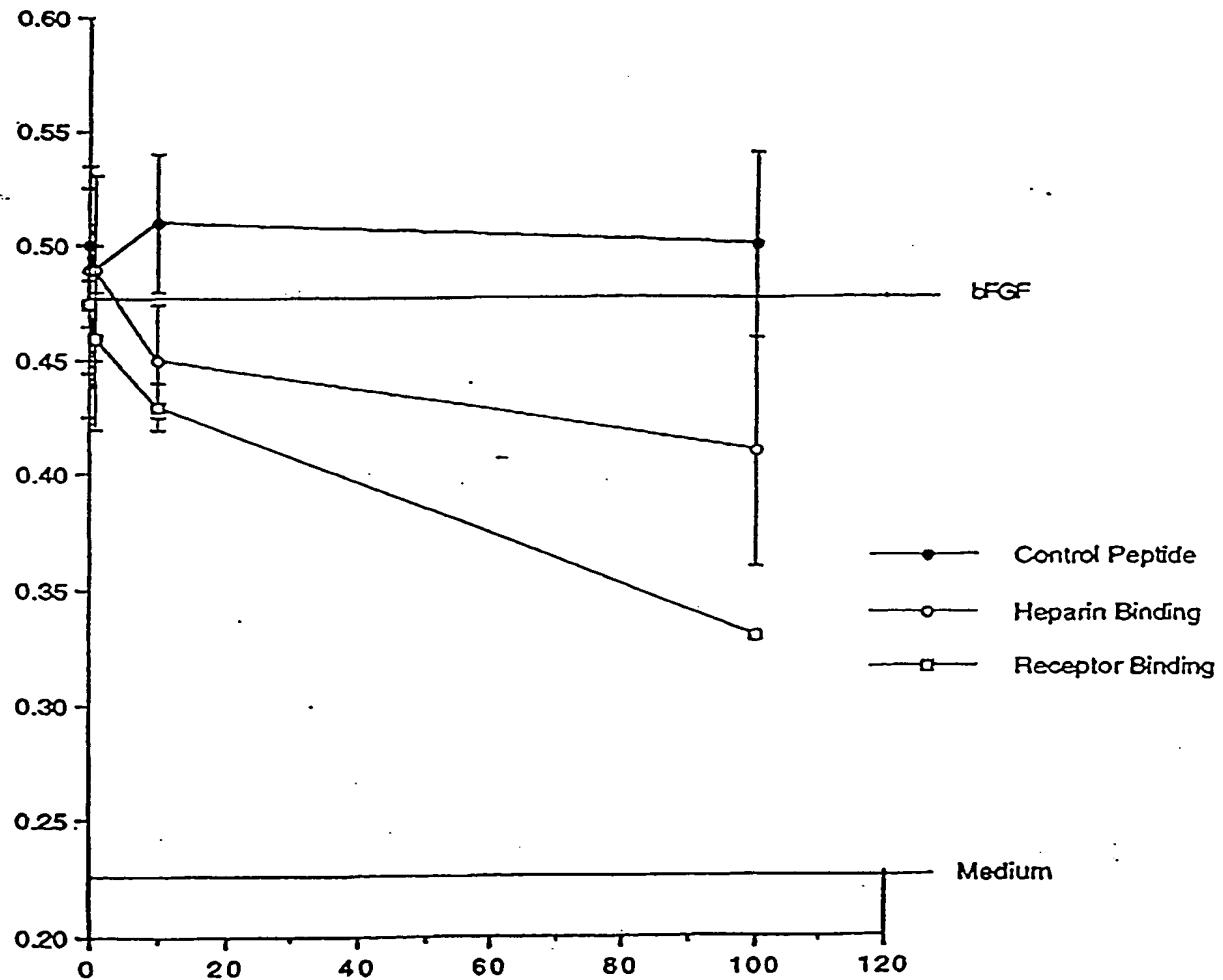


Figure 8

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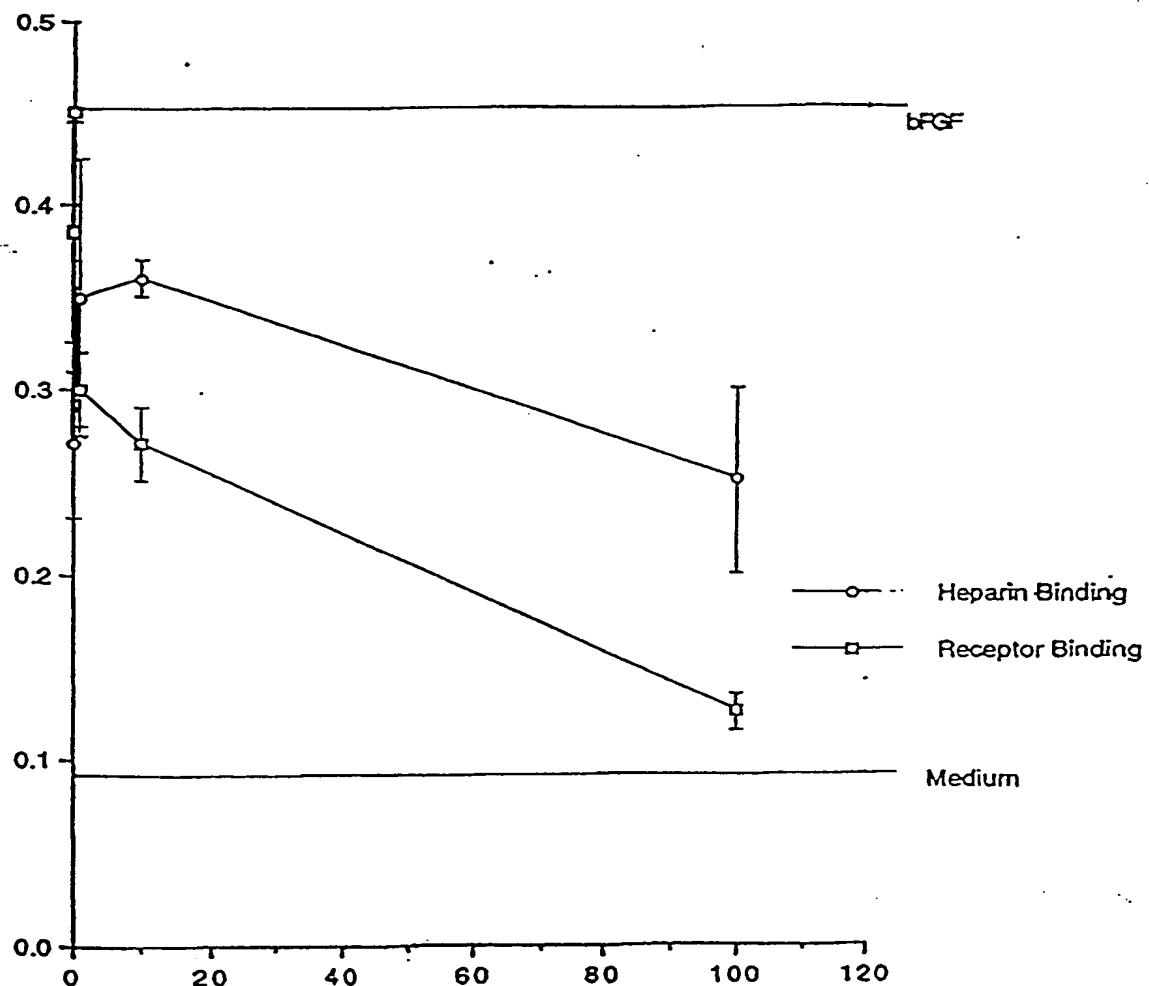


Figure 9

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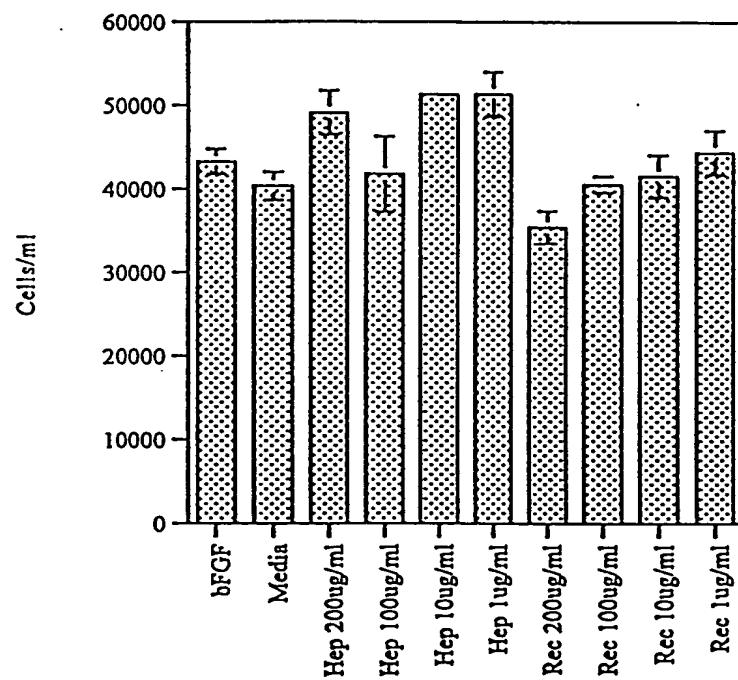


Figure 10

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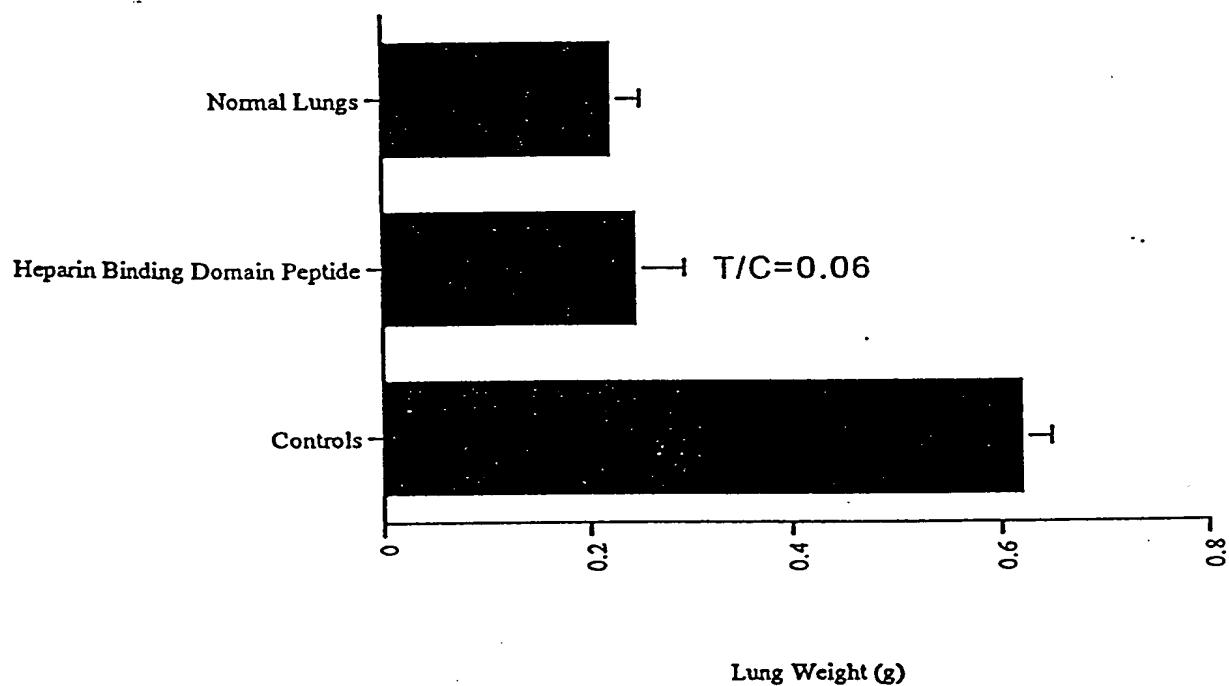


Figure 11

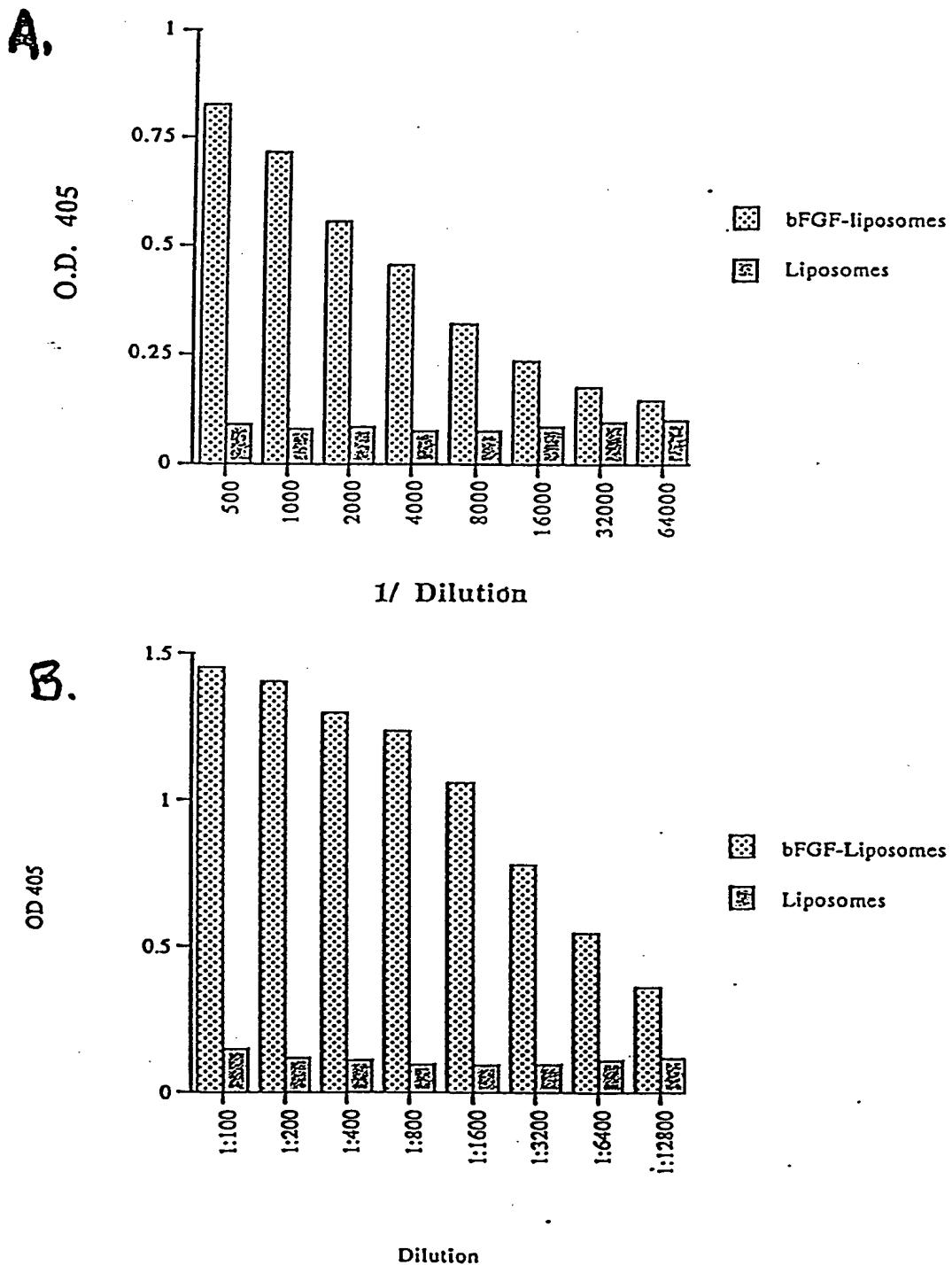


Figure 12

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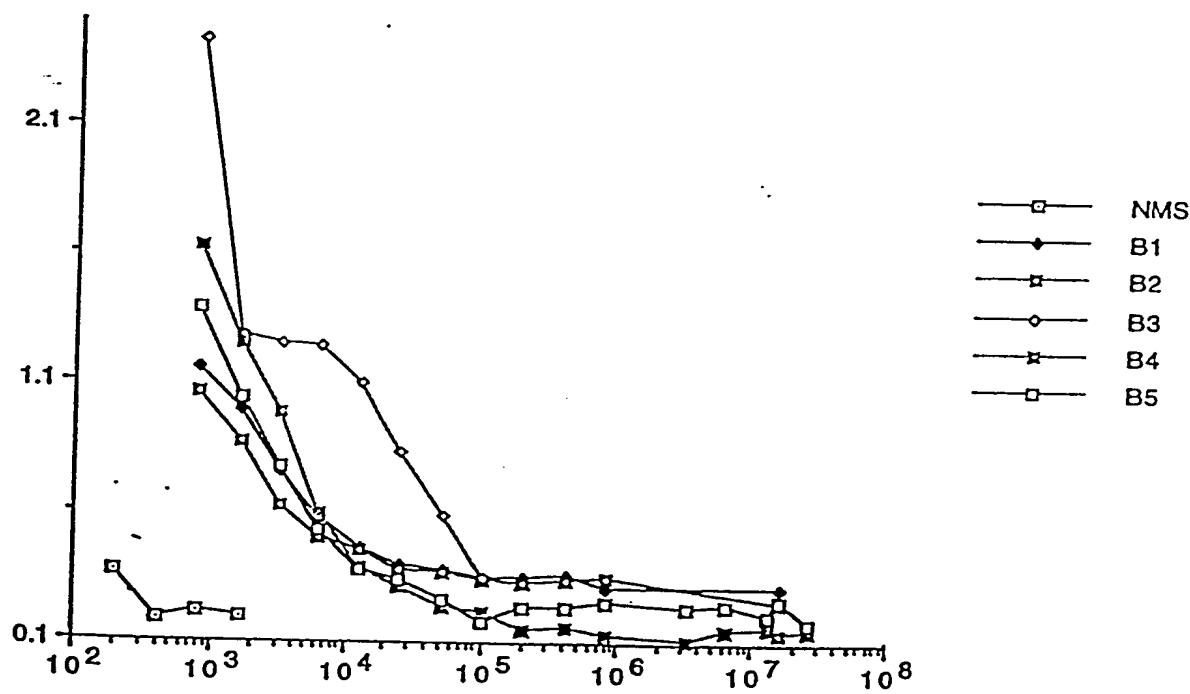


Figure 13

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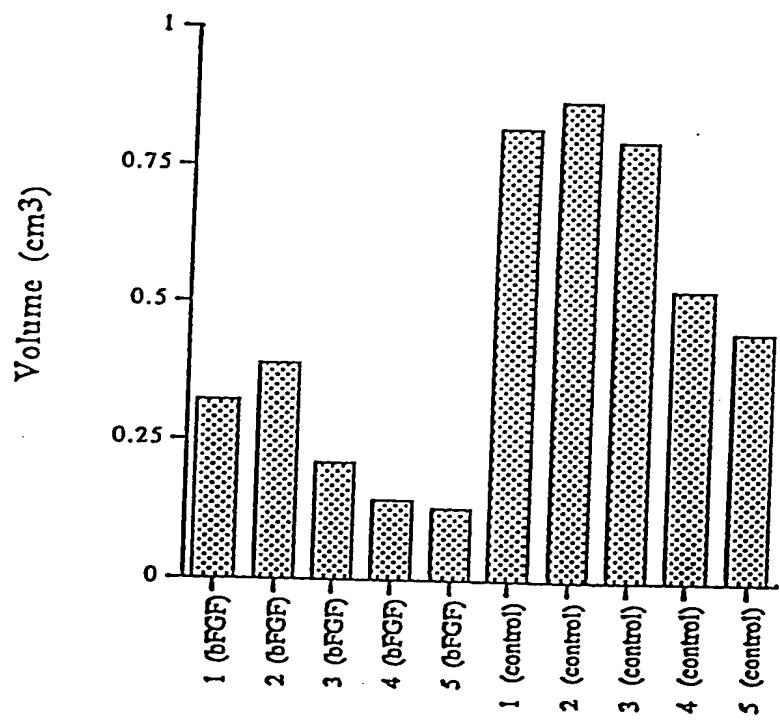


Figure 14

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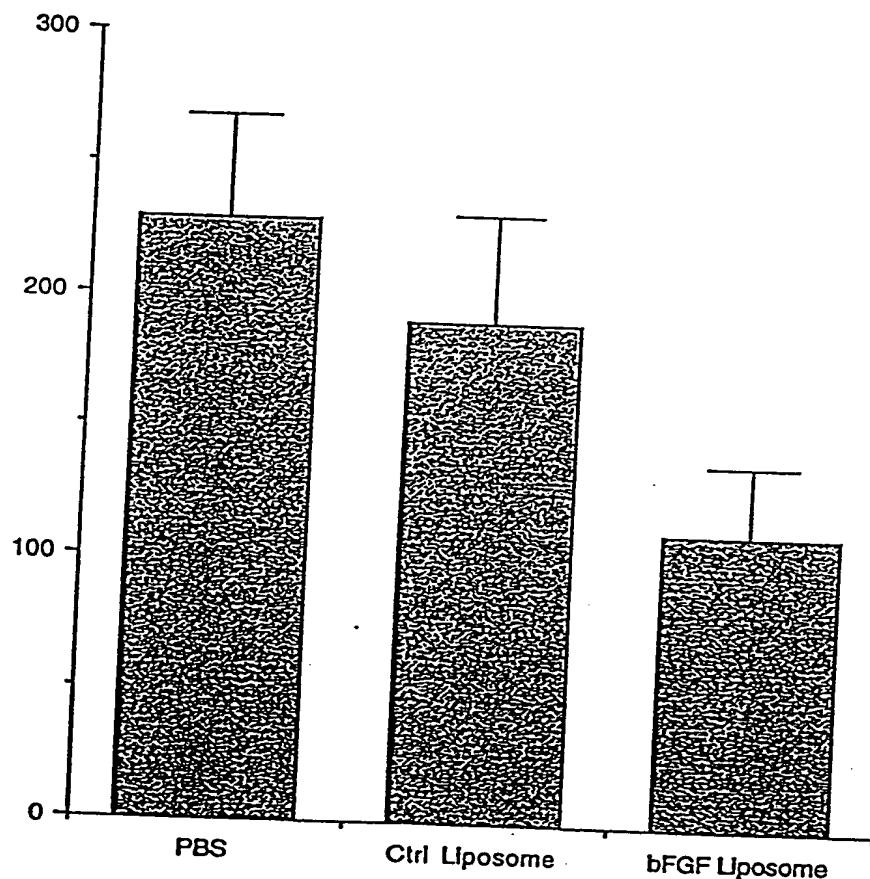
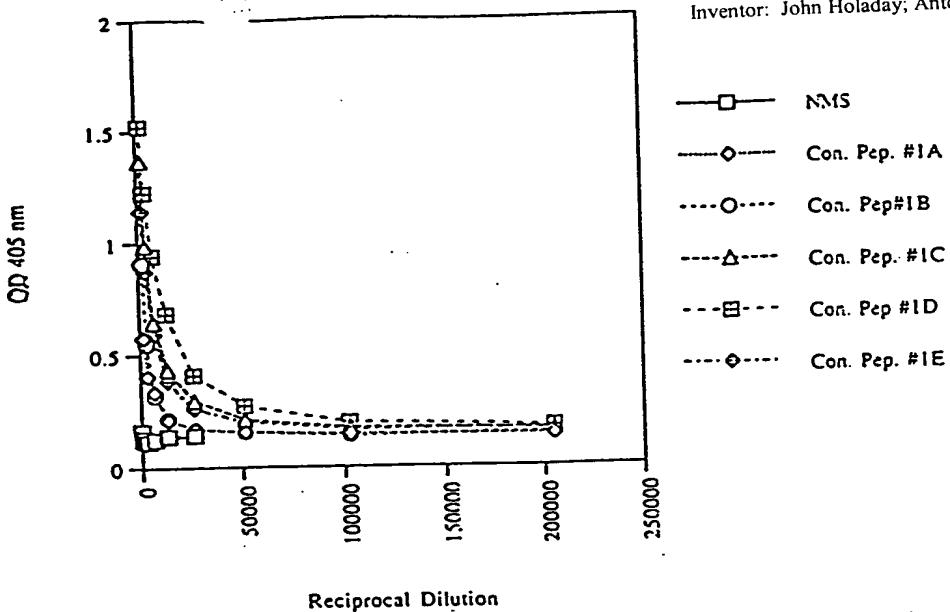
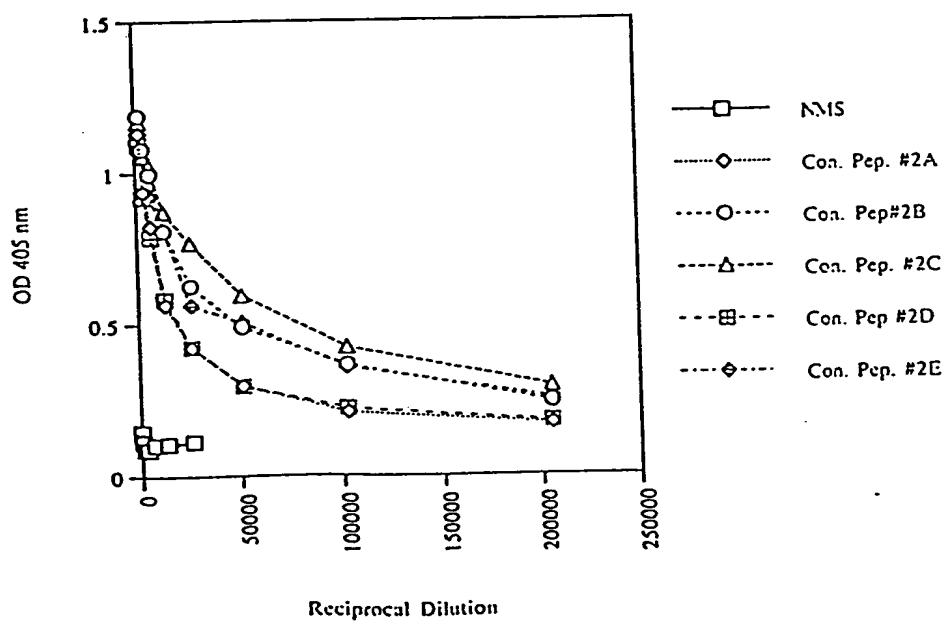


Figure 15

A



B



C

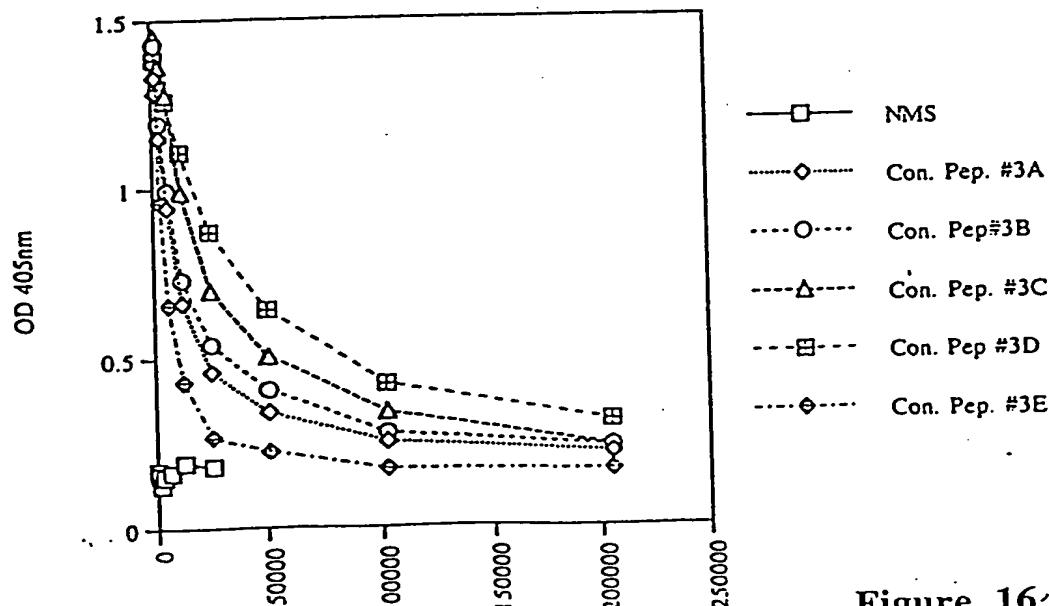
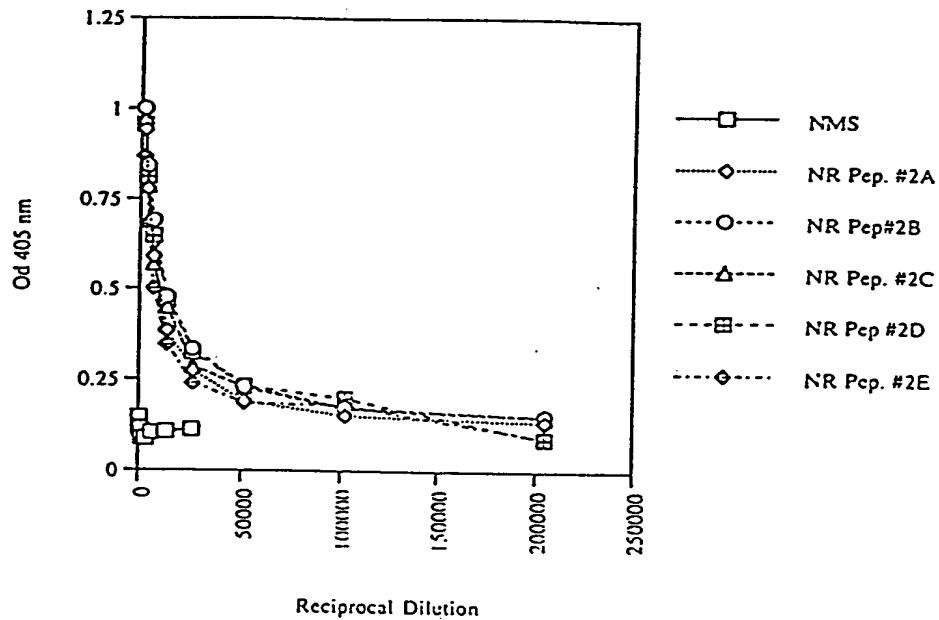


Figure 16

A



B

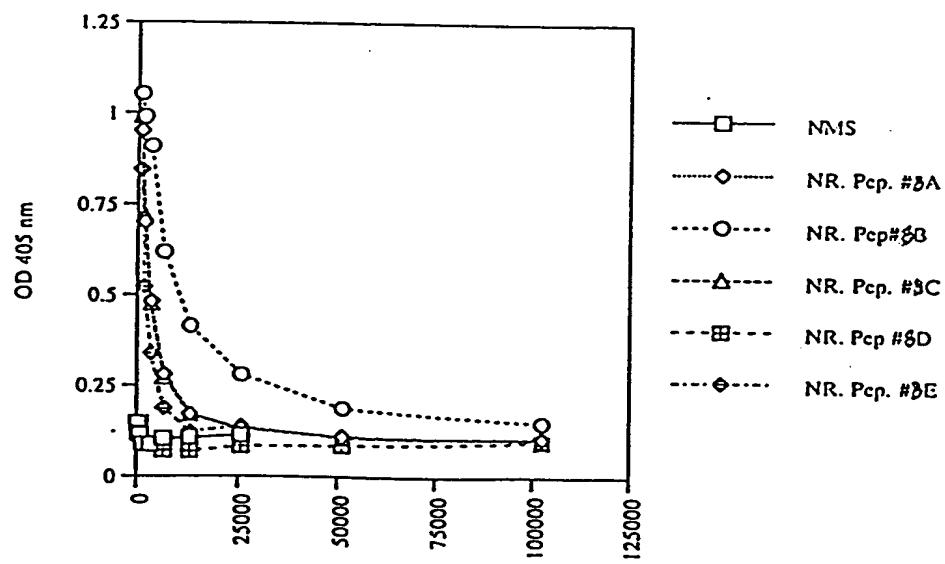


Figure 17

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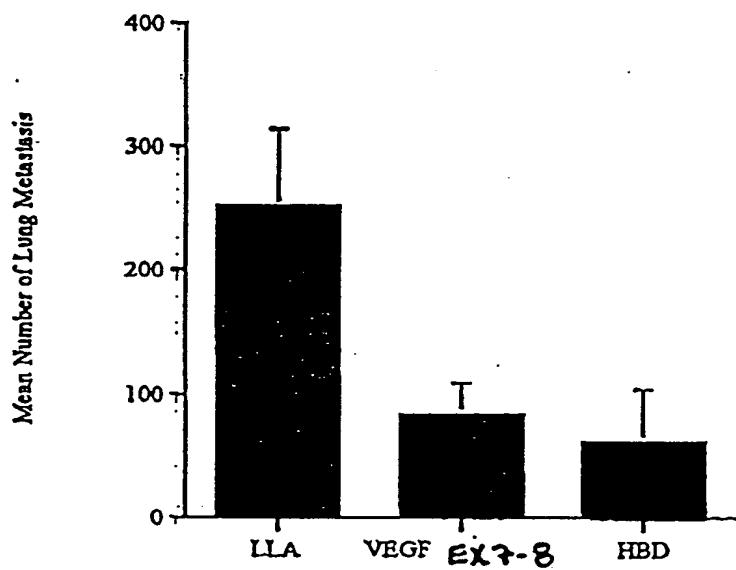


Figure 18

Figure 19: Overview

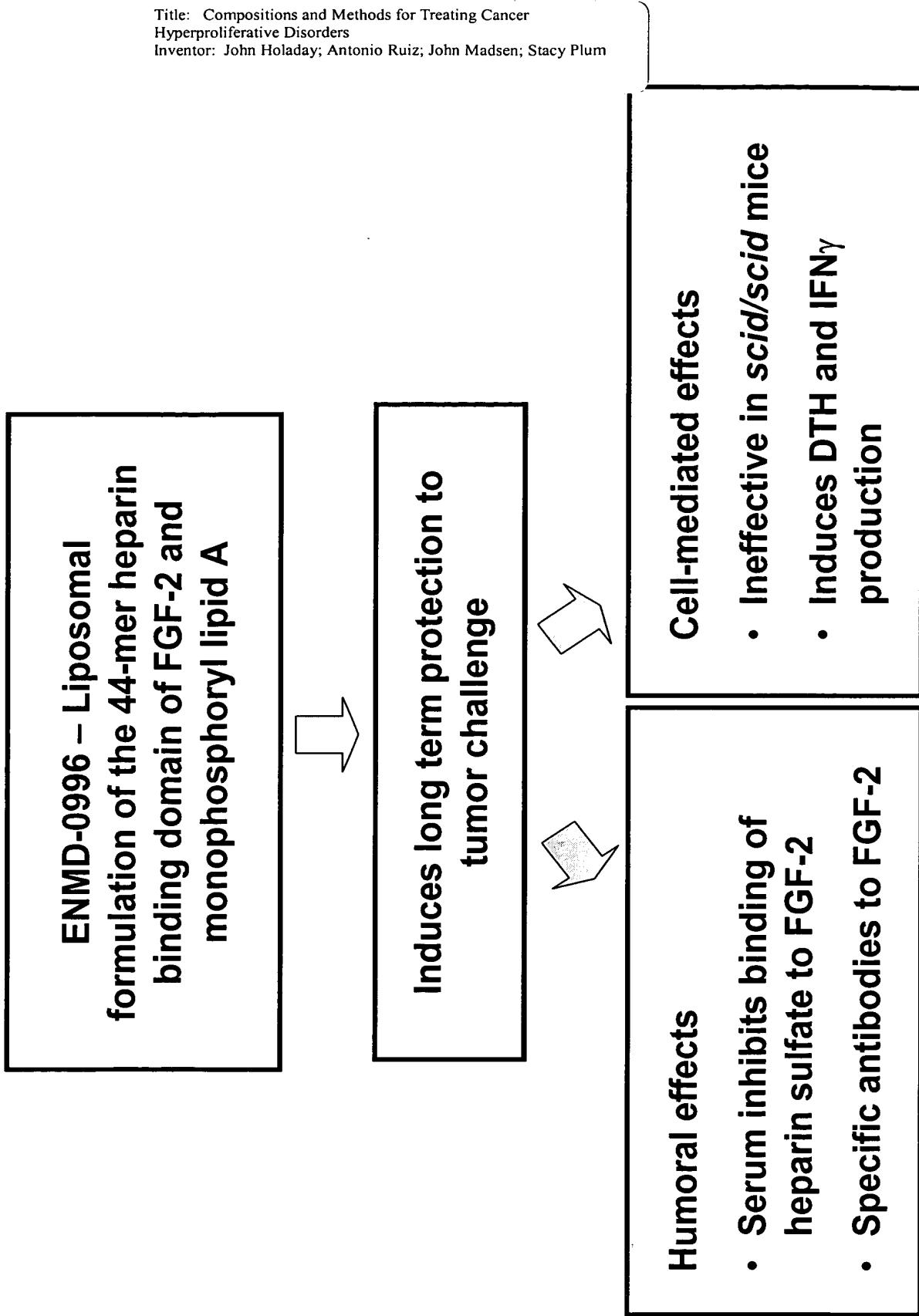


Figure 20: ENMD-0996 induces long-term protection against tumor challenge in mice

Challenge date	Treatment	Activity in B16BL6	
		Mean number of metastasis	T/C
2 Weeks	ENMD-0996	47 ± 37	0.20
	Control	234 ± 82	1.00
3 Months	ENMD-0996	61 ± 42	0.24
	Control	252 ± 62	1.00

Figure 21: Sera from ENMD-0996-treated Mice Inhibit the Binding of Heparin Sulfate to FGF-2

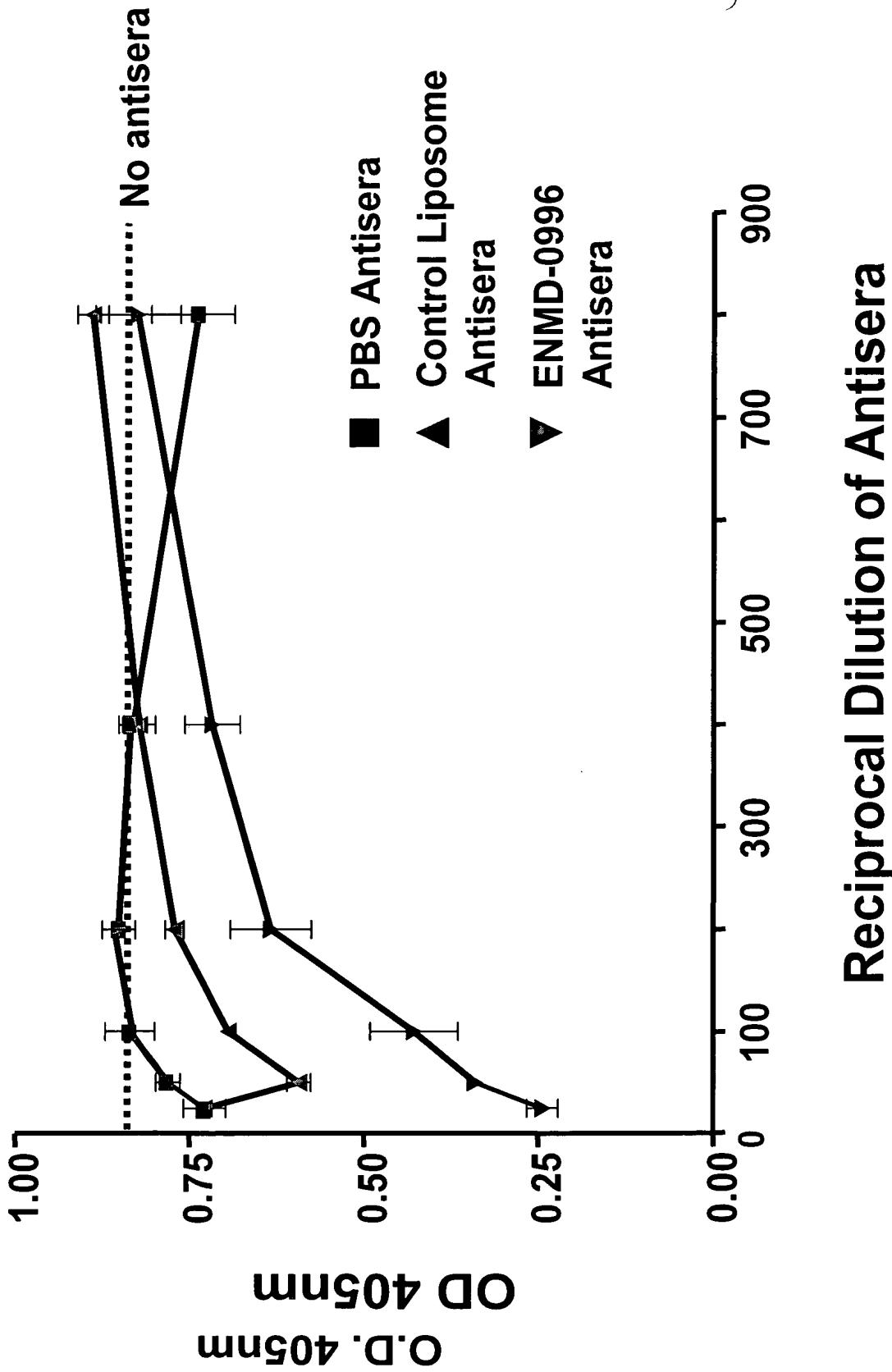


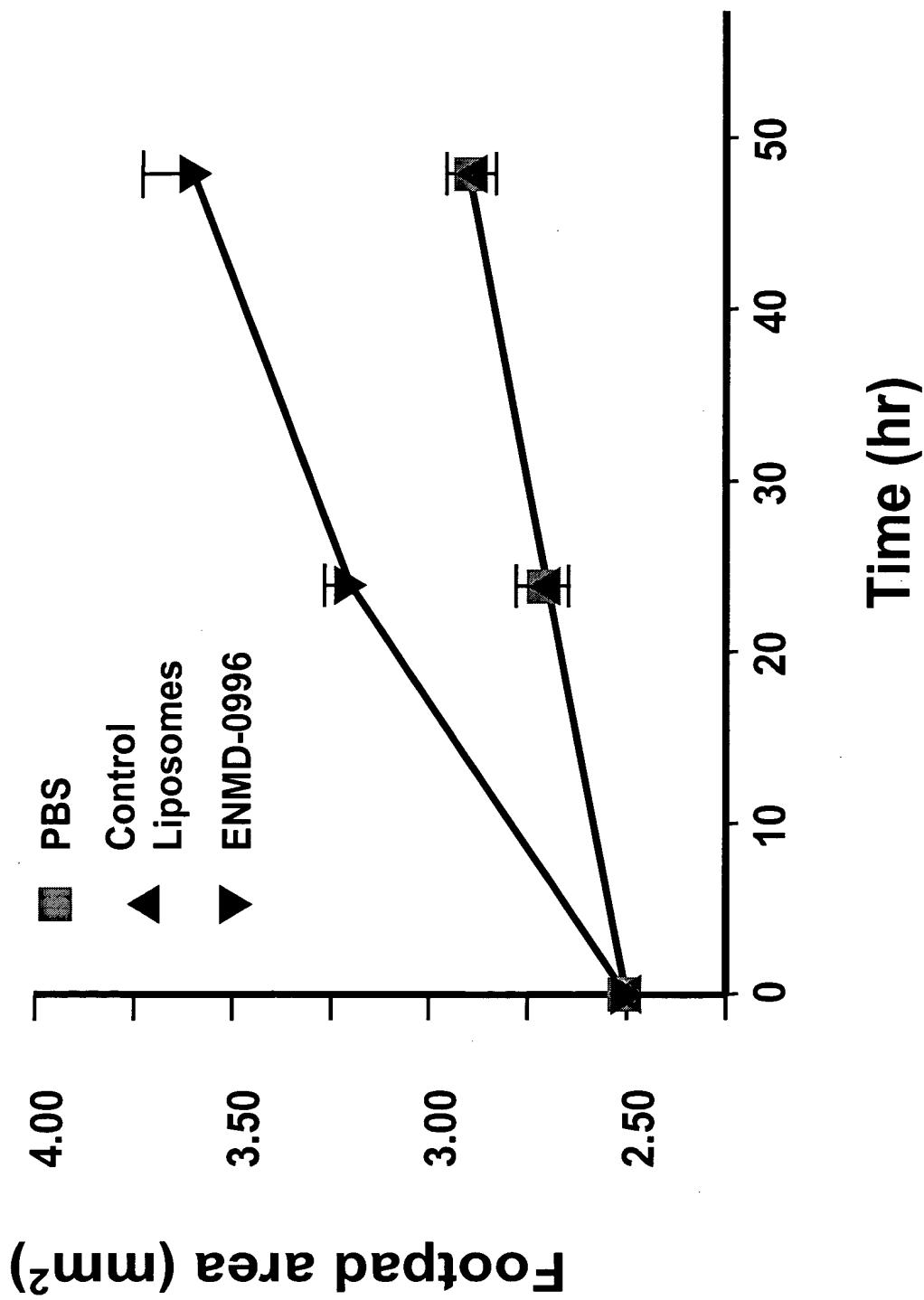
Figure 22. The antibody response in mice treated with ENMD-0996 is specific for FGF-2

Antigen	Antibody Titer
Basic FGF/FGF-2	1:5000
Acidic FGF/FGF-1	< 1:100
FGF-3	< 1:100
FGF-4	< 1:100
FGF-5	< 1:100
FGF-6	< 1:100
FGF-7	< 1:100
FGF-9	< 1:100
FGF-10	< 1:100
FGF-17	< 1:100
VEGF	< 1:100
TFPI	< 1:100

Figure 23: ENMD-0996 does not inhibit metastasis in mice lacking functional B- and T-cells

Treatment	Mice	Mean number of Metastasis (+ SD)	T/C
Control Liposomes	C57BL/6	105 ± 16	1.00
	C57BL/6	146 ± 29	1.40
	C57BL/6	37 ± 5	0.35
ENMD-0996	C57BL/6 scID/scID	103 ± 18	1.00
	C57BL/6 scID/scID	100 ± 22	0.97
	C57BL/6 scID/scID	86 ± 28	0.83

Figure 24: Mice Treated with ENMD-0996 Develop a Delayed Type Hypersensitivity to Subsequent Challenge with FGF-2



**Figure 25. Treatment of Mice with ENMD-0996
Results in the Production of IFN_γ Production**

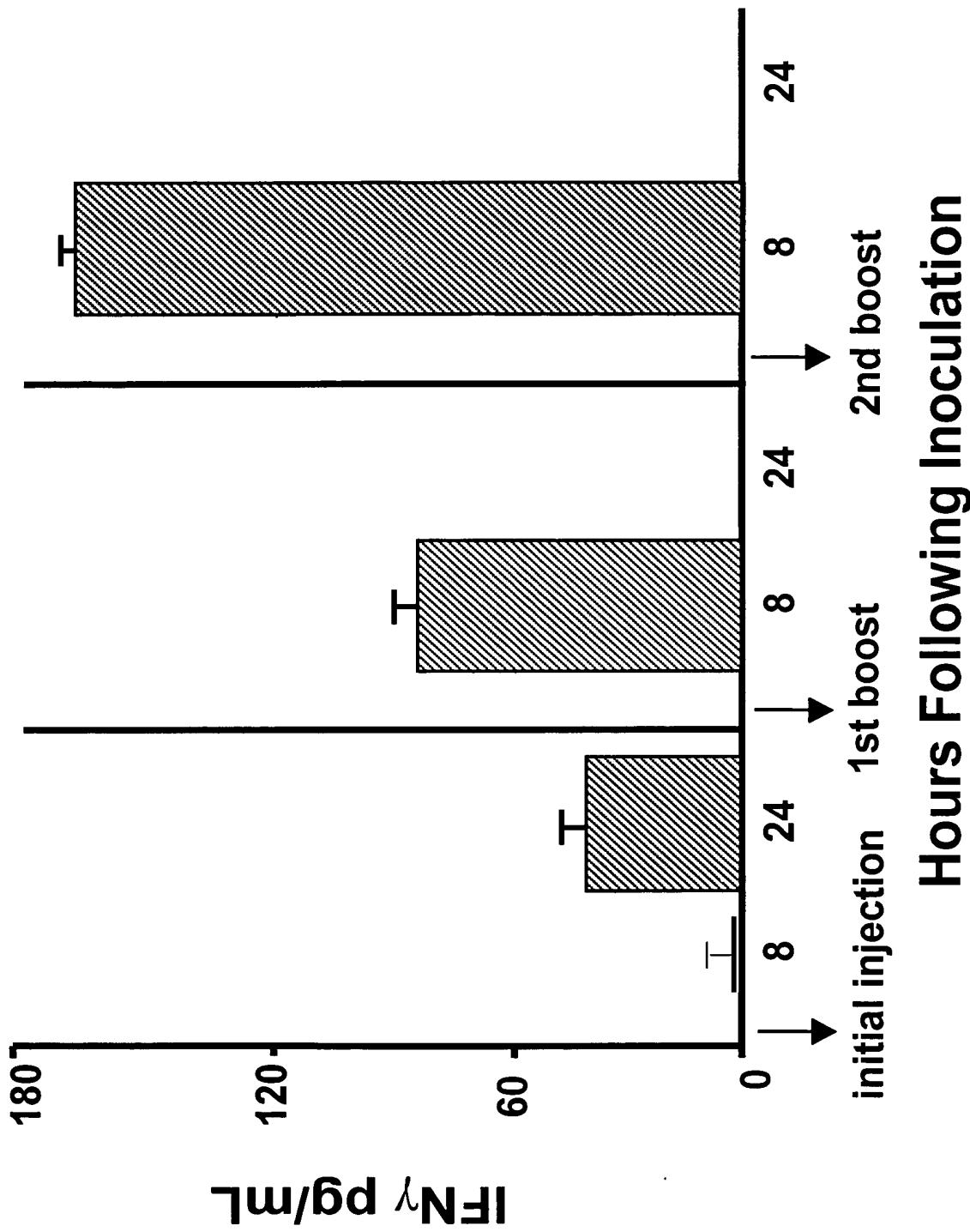


Figure 26: Splenocytes from mice treated with ENMD-0996 produce IFN_γ in response to FGF-2 or HBD

